

Package ‘scGOclust’

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Type Package

Title Measuring Cell Type Similarity with Gene Ontology in Single-Cell RNA-Seq

Version 0.2.1

Description Traditional methods for analyzing single cell RNA-seq datasets focus solely on gene expression, but this package introduces a novel approach that goes beyond this limitation. Using Gene Ontology terms as features, the package allows for the functional profile of cell populations, and comparison within and between datasets from the same or different species. Our approach enables the discovery of previously unrecognized functional similarities and differences between cell types and has demonstrated success in identifying cell types' functional correspondence even between evolutionarily distant species.

URL <https://github.com/Papatheodorou-Group/scGOclust>

BugReports <https://github.com/Papatheodorou-Group/scGOclust/issues>

License GPL (>= 3)

Encoding UTF-8

LazyData true

LazyDataCompression bzip2

RoxygenNote 7.2.3

Imports limma, Seurat(>= 5.0.0), biomaRt, dplyr, magrittr, stats, tibble, tidyr, Matrix, utils, networkD3, slanter

Suggests knitr, devtools, pheatmap, rmarkdown, httr

VignetteBuilder knitr

Depends R (>= 2.10)

NeedsCompilation no

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analyzeGOSeurat	<i>standard seurat analysis on GO_seurat object</i>
-----------------	---

Description

standard seurat analysis on GO_seurat object

Usage

```
analyzeGOSeurat(
  go_seurat_obj,
  cell_type_col,
  norm_log1p = TRUE,
  scale.factor = 10000,
  nfeatures = 2000,
  cluster_res = 1,
  min.dist = 0.3,
  ...
)
```

Arguments

go_seurat_obj	go seurat object created by makeGOSeurat
cell_type_col	column name in mera.data storing cell type classes
norm_log1p	whether or not to perform data normalisation and log1p transformation, default TRUE
scale.factor	param for Seurat NormalizeData

nfeatures	param for Seurat FindVariableFeatures
cluster_res	resolution for Seurat FindClusters
min.dist	param for Seurat RunUMAP
...	additional params for all Seurat functions involved in this function

Value

standard analyzed GO seurat object until UMAP

Examples

```
library(scG0clust)
library(httr)
httr::set_config(httr::config(ssl_verifypeer = FALSE))
data(mmu_tbl)
data(mmu_subset)
go_seurat_obj = makeGOSeurat(ensembl_to_GO = mmu_tbl,
  seurat_obj = mmu_subset,
  feature_type = "external_gene_name")

analyzeGOSeurat(go_seurat_obj = go_seurat_obj, cell_type_col = "cell_type_annotation")
```

cellTypeGOCorr	<i>calculate correlation between cell types represented by scaled GO, per-species</i>
----------------	---

Description

calculate correlation between cell types represented by scaled GO, per-species

Usage

```
cellTypeGOCorr(cell_type_go, corr_method = "pearson")
```

Arguments

cell_type_go	cell type GO table calculated via getCellTypeGO
corr_method	correlation method, choose among "pearson", "kendall", "spearman", default 'pearson'

Value

a dataframe with correlation between cell types

Examples

```

library(scGOclust)
library(httr)
httr::set_config(httr::config(ssl_verifypeer = FALSE))
data(mmu_tbl)
data(mmu_subset)
go_seurat_obj = makeGOSeurat(ensembl_to_GO = mmu_tbl,
  seurat_obj = mmu_subset,
  feature_type = "external_gene_name")

cell_type_go = getCellTypeGO(go_seurat_obj = go_seurat_obj, cell_type_co = "cell_type_annotation")

cellTypeGOCorr(cell_type_go = cell_type_go, corr_method = "pearson")

```

crossSpeciesCellTypeGOCorr

calculate cross-species correlation between cell types represented by scaled GO

Description

calculate cross-species correlation between cell types represented by scaled GO

Usage

```

crossSpeciesCellTypeGOCorr(
  species_1,
  species_2,
  cell_type_go_sp1,
  cell_type_go_sp2,
  corr_method = "pearson"
)

```

Arguments

species_1	name of species one
species_2	name of species two
cell_type_go_sp1	cell type GO table of species one calculated via getCellTypeGO
cell_type_go_sp2	cell type GO table of species two calculated via getCellTypeGO
corr_method	correlation method, choose among "pearson", "kendall", "spearman", default 'pearson'

Value

correlation between cell types

Examples

```
library(scGOclust)
library(httr)
httr::set_config(httr::config(ssl_verifypeer = FALSE))
data(mmu_tbl)
data(mmu_subset)
data(dme_tbl)
data(dme_subset)
mmu_go_obj = makeGOSeurat(ensembl_to_GO = mmu_tbl,
  seurat_obj = mmu_subset,
  feature_type = "external_gene_name")
dme_go_obj = makeGOSeurat(ensembl_to_GO = dme_tbl,
  seurat_obj = dme_subset,
  feature_type = "external_gene_name")

mmu_cell_type_go = getCellTypeGO(go_seurat_obj = mmu_go_obj, cell_type_co = "cell_type_annotation")
dme_cell_type_go = getCellTypeGO(go_seurat_obj = dme_go_obj, cell_type_co = "annotation")

crossSpeciesCellTypeGOCorr(species_1 = 'mmusculus',
  species_2 = 'dmelanogaster',
  cell_type_go_sp1 = mmu_cell_type_go,
  cell_type_go_sp2 = dme_cell_type_go)
```

dme_subset	<i>Drosophila gut scRNA-seq data, 10X Chromium Subset to 45 cells per cell type as an example data</i>
------------	--

Description

Drosophila gut scRNA-seq data, 10X Chromium Subset to 45 cells per cell type as an example data

Usage

```
dme_subset
```

Format

a 'Seurat' object

Source

<<https://flycellatlas.org/>>

dme_tbl	<i>Drosophila EMSEMBL gene and GO annotation, subset to genes present in 'dme_subset'</i>
---------	---

Description

Drosophila EMSEMBL gene and GO annotation, subset to genes present in 'dme_subset'

Usage

```
dme_tbl
```

Format

a 'data.frame' object

Source

<<http://www.ensembl.org/>>

ensemblToGo	<i>get requested ensembl ID to GO mapping table</i>
-------------	---

Description

get requested ensembl ID to GO mapping table

Usage

```
ensemblToGo(
  species,
  GO_type = "biological_process",
  GO_linkage_type = c("standard"),
  ...
)
```

Arguments

species	species name matching ensembl biomaRt naming, such as hsapiens, mmusculus
GO_type	GO term type, choose among 'biological_process', 'molecular_function', 'cellular_component', default 'biological_process'

GO_linkage_type
 GO annotation evidence codes to include. Default is 'standard', which means only including manually checked records (excluding IEA) and excluding those inferred from gene expression experiments to maximally suffice the species expression independence assumption. 'Stringent' means only including those with experimental evidence, also not from gene expression experiment, or from manual curation with evidence (excluding those from mass-annotation pipelines). Choose among 'experimental', 'phylogenetic', 'computational', 'author', 'curator', 'electronic', 'standard', 'stringent'

... additional params for useEnsembl function called in this function

Value

a table with ensembl to GO terms mapping including requested linkage type

Examples

```
library(scGOclust)
library(httr)
httr::set_config(httr::config(ssl_verifypeer = FALSE))
ensemblToGo("mmusculus", GO_type = "biological_process", GO_linkage_type = 'stringent')
```

getCellTypeGO	<i>get per cell type average scaled vector of GO terms</i>
---------------	--

Description

get per cell type average scaled vector of GO terms

Usage

```
getCellTypeGO(go_seurat_obj, cell_type_col, norm_log1p = TRUE)
```

Arguments

go_seurat_obj go seurat object created by makeGOSeurat
 cell_type_col column name in mera.data storing cell type classes
 norm_log1p whether or not to perform data normalisation and log1p transformation, default TRUE

Value

a table of scaled GO representation per cell type (averaged)

Examples

```

library(scG0clust)
library(httr)
httr::set_config(httr::config(ssl_verifypeer = FALSE))
data(mmu_tbl)
data(mmu_subset)
go_seurat_obj = makeGOSeurat(ensembl_to_GO = mmu_tbl,
  seurat_obj = mmu_subset,
  feature_type = "external_gene_name")
getCellTypeGO(go_seurat_obj = go_seurat_obj, cell_type_co = "cell_type_annotation")

```

```

getCellTypeSharedGO  get shared up and down regulated GO terms for all pairs of cell types

```

Description

get shared up and down regulated GO terms for all pairs of cell types

Usage

```

getCellTypeSharedGO(
  species_1,
  species_2,
  analyzed_go_seurat_sp1,
  analyzed_go_seurat_sp2,
  cell_type_col_sp1,
  cell_type_col_sp2,
  layer_use = "data",
  p_val_threshold = 0.01
)

```

Arguments

```

species_1      name of species one
species_2      name of species two
analyzed_go_seurat_sp1
                analyzed GO seurat object of species one
analyzed_go_seurat_sp2
                analyzed GO seurat object of species two
cell_type_col_sp1
                cell type column name for species 1 data
cell_type_col_sp2
                cell type column name for species 2 data

```


layer_use layer to use for marker computation, default 'data' which after NormalizeData will be log1p normalized data.

p_val_threshold p value threshold for selecting DEG (p_adjust)

Value

a list with sp1 raw, sp2 raw and shared, significant up and down regulated GO terms per cell type (pair)

Examples

```
library(scGOclust)
library(httr)
httr::set_config(httr::config(ssl_verifypeer = FALSE))
data(mmu_tbl)
data(mmu_subset)
data(dme_tbl)
data(dme_subset)

mmu_go_obj = makeGOSeurat(ensembl_to_GO = mmu_tbl,
  seurat_obj = mmu_subset,
  feature_type = "external_gene_name")
dme_go_obj = makeGOSeurat(ensembl_to_GO = dme_tbl,
  seurat_obj = dme_subset,
  feature_type = "external_gene_name")

mmu_go_obj_analyzed = analyzeGOSeurat(mmu_go_obj, "cell_type_annotation")
dme_go_obj_analyzed = analyzeGOSeurat(dme_go_obj, "annotation")

getCellTypeSharedGO(species_1 = 'mmusculus',
  species_2 = 'dmelanogaster',
  analyzed_go_seurat_sp1 = mmu_go_obj_analyzed,
  analyzed_go_seurat_sp2 = dme_go_obj_analyzed,
  cell_type_col_sp1 = 'cell_type_annotation',
  cell_type_col_sp2 = 'annotation',
  layer_use = "data",
  p_val_threshold = 0.01)
```

getCellTypeSharedTerms

query co-up and co-down regulated GO terms from certain cell type pairs

Description

query co-up and co-down regulated GO terms from certain cell type pairs

Usage

```
getCellTypeSharedTerms(
  shared_go,
  cell_type_sp1,
  cell_type_sp2,
  return_full = FALSE,
  arrange_avg_log2FC = TRUE
)
```

Arguments

```
shared_go      cell type shared GO table from getCellTypeSharedGO
cell_type_sp1  cell type from sp1 to query
cell_type_sp2  cell type from sp2 to query
return_full    if return also pvals and logfc info, default FALSE
arrange_avg_log2FC
                arrange result by decreasing mean avg_log2FC, default TRUE
```

Value

a dataframe displaying co-up or co-down regulated GO terms for the queried cell type pair

Examples

```
library(scGOclust)
library(httr)
httr::set_config(httr::config(ssl_verifypeer = FALSE))
data(mmu_tbl)
data(mmu_subset)
data(dme_tbl)
data(dme_subset)

mmu_go_obj = makeGOSeurat(ensembl_to_GO = mmu_tbl,
  seurat_obj = mmu_subset,
  feature_type = "external_gene_name")
dme_go_obj = makeGOSeurat(ensembl_to_GO = dme_tbl,
  seurat_obj = dme_subset,
  feature_type = "external_gene_name")

mmu_go_obj_analyzed = analyzeGOSeurat(mmu_go_obj, "cell_type_annotation")
dme_go_obj_analyzed = analyzeGOSeurat(dme_go_obj, "annotation")

shared_go = getCellTypeSharedGO(species_1 = 'mmusculus',
  species_2 = 'dmelanogaster',
  analyzed_go_seurat_sp1 = mmu_go_obj_analyzed,
  analyzed_go_seurat_sp2 = dme_go_obj_analyzed,
  cell_type_col_sp1 = 'cell_type_annotation',
  cell_type_col_sp2 = 'annotation',
```

```
layer_use = "data",
p_val_threhold = 0.01)
```

```
getCellTypeSharedTerms(shared_go = shared_go,
cell_type_sp1 = 'intestine_Enterоendocrine cell',
cell_type_sp2 = 'enterоendocrine cell',
return_full = FALSE)
```

globalvariables	<i>record some global variables: pre-defined column name in biomaRt query and markers</i>
-----------------	---

Description

record some global variables: pre-defined column name in biomaRt query and markers

makeGOSeurat	<i>create a seurat object with GO terms</i>
--------------	---

Description

create a seurat object with GO terms

Usage

```
makeGOSeurat(ensembl_to_GO, seurat_obj, feature_type = "ensembl_gene_id")
```

Arguments

ensembl_to_GO	ensembl_to_go mapping table from function ensemblToGo
seurat_obj	count matrix with genes to cells
feature_type	feature type of count matrix, choose from ensembl_gene_id, external_gene_name, default ensembl_gene_id

Value

a seurat object with GO terms as features

Examples

```
library(scG0clust)
library(httr)
httr::set_config(httr::config(ssl_verifypeer = FALSE))
data(mmu_tbl)
data(mmu_subset)
makeG0Seurat(ensembl_to_G0 = mmu_tbl,
  seurat_obj = mmu_subset,
  feature_type = "external_gene_name")
```

mmu_subset	<i>Mouse stomach and intestine scRNA-seq data, microwell-seq Subset to 50 cells per cell type as an example data</i>
------------	--

Description

Mouse stomach and intestine scRNA-seq data, microwell-seq Subset to 50 cells per cell type as an example data

Usage

```
mmu_subset
```

Format

a 'Seurat' object

Source

<<https://bis.zju.edu.cn/MCA/>>

mmu_tbl	<i>Mouse EMSEMBL gene and GO annotation, subset to genes present in 'mmu_subset'</i>
---------	--

Description

Mouse EMSEMBL gene and GO annotation, subset to genes present in 'mmu_subset'

Usage

```
mmu_tbl
```

Format

a 'data.frame' object

Source

<<http://www.ensembl.org/>>

plotCellTypeCorrHeatmap

plot clustered heatmap for cell type corr

Description

plot clustered heatmap for cell type corr

Usage

```
plotCellTypeCorrHeatmap(corr_matrix, scale = NA, ...)
```

Arguments

corr_matrix	correlation matrix from cellTypeGOCorr or crossSpeciesCellTypeGOCorr
scale	scale value by column, row, or default no scaling (NA)
...	params to pass to slanter::sheatmap

Value

a sheatmap heatmap

Examples

```
library(scGOclust)
library(httr)
httr::set_config(httr::config(ssl_verifypeer = FALSE))
data(mmu_tbl)
data(mmu_subset)

go_seurat_obj = makeGOSeurat(ensembl_to_GO = mmu_tbl,
  seurat_obj = mmu_subset,
  feature_type = "external_gene_name")

cell_type_go = getCellTypeGO(go_seurat_obj = go_seurat_obj, cell_type_co = "cell_type_annotation")

corr_matrix = cellTypeGOCorr(cell_type_go = cell_type_go, corr_method = "pearson")

plotCellTypeCorrHeatmap(corr_matrix = corr_matrix, scale = "column")
```

plotCellTypeSankey *plot Sankey diagram for cell type links above a certain threshold*

Description

plot Sankey diagram for cell type links above a certain threshold

Usage

```
plotCellTypeSankey(corr_matrix, corr_threshold = 0.1, ...)
```

Arguments

corr_matrix	cell type corr matrix from crossSpeciesCellTypeGOCorr
corr_threshold	minimum corr value for positively related cell types, default 0.6
...	additional params for sankeyNetwork

Value

a Sankey plot showing related cell types

Examples

```
library(scG0clust)
library(httr)
httr::set_config(httr::config(ssl_verifypeer = FALSE))
data(mmu_tbl)
data(mmu_subset)
go_seurat_obj = makeG0Seurat(ensembl_to_GO = mmu_tbl,
  seurat_obj = mmu_subset,
  feature_type = "external_gene_name")

cell_type_go = getCellTypeG0(go_seurat_obj = go_seurat_obj, cell_type_co = "cell_type_annotation")
corr_matrix = cellTypeGOCorr(cell_type_go = cell_type_go, corr_method = "pearson")

plotCellTypeSankey(corr_matrix = corr_matrix, 0.1)
```

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